Outcomes of Weight Loss During Induction Therapy During Childhood ALL

JIT HEALTH SCIENCE CENTER

Pediatric Research Program

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INTRODUCTION

Acute lymphoblastic leukemia (ALL) is the most common childhood cancer. Therapy involves multiple phases such as induction, consolidation, interim maintenance, delayed intensification and then a longer term maintenance phase. Pediatric patients with ALL and lymphoblastic lymphoma (LLy) often experience significant weight gain (≥5% weight gain) during remission induction therapy. However, some experience significant weight loss (≥5% weight loss) instead.

There are several factors associated with a decrease in body mass index (BMI) during therapy: high/very high risk stratification, being overweight/obese at diagnosis, trisomy 21, and hyperglycemia. It is important to monitor malnutrition due the potential outcomes of decreased survival, infection risk, and rapid loss of lean body mass.

The purpose of this study is to determine whether significant weight loss during induction therapy decreased the likelihood of reaching end of therapy (EOT) or increased risk of death or relapse in these patients.

METHODS

- Retrospective chart review of pediatric patients diagnosed with ALL or LLy at Cook Children's Medical Center (CCMC) from 1/1/11 to 3/31/17.
- Inclusion criteria:
 - ≥2 years of age at diagnosis and <20 years of age by day 1 of consolidation
- Exclusion criteria:
 - If induction was not completed at CCMC
 - If patients were not treated according to the following COG protocols: AALL0331, AALL0232, AALL0434, AALL0932, AALL1131, AALL1231, AALL0622, AALL1122
- Percent weight change from diagnosis to end of induction was grouped: loss (≥5% weight loss), gain (≥5% weight gain), and steady (<5% weight loss/gain).
- Data was collected regarding the following outcomes (if applicable):
 - Death
 - Relapse
 - Whether end of therapy (EOT) was reached
- To examine outcomes of weight loss, logistic regression was used for reaching EOT and Cox regression for death and relapse.

RESULTS

- There were 187 patients included in the study. Weight-change categories:
 - 17% loss
 - 39% steady
 - 45% gain
- Outcomes Data:
 - Relapse: 22 patients (12%)
 - EOT Not Reached: 18 patients (10%)
 - Death: 10 patients (5%)
- Compared to patients in the steady category, patients who lost weight were significantly less likely to reach EOT.
- Though nonsignificant, the odds ratio demonstrated increased risk of death and relapse, but this did not reach significance.
- Patients in the steady and gain groups did not significantly differ in any outcomes of EOT, death, or relapse (p>0.05).

	Loss vs. Steady	Gain vs. Steady	
Variable	OR/HR (95% CI)	p OR/HR (95% CI)	ŗ
End of Therapy ^a	0.31 (0.16, 0.63) 0.002	1 1.98 (0.95, 4.11) 0.0	67
Death ^b	3.67 (0.81, 16.52) 0.090	0 0.81 (0.16, 4.03) 0.7	'99
Relapseb	1.83 (0.60, 5.60) 0.292	1 0.93 (0.36, 2.40) 0.8	76
Event ^b	1.93 (0.68, 5.43) 0.21	4 0.82 (0.33, 2.07) 0.6	76

OR/HR, odds or hazard ratio; 95% CI, 95% confidence interval; P, P-value; Dx, diagnosis.

^aLogistic regression ^bCox regression

CONCLUSION

This research demonstrates that patients who experience malnutrition during induction therapy may be at greater risk for not completing therapy. Those in the loss group had higher mortality rates (13%) than did those in the steady or gain (4%) groups; and a larger sample may have produced a significant effect for death. Therefore, further study is necessary to determine if weight loss during induction therapy should be considered for additional nutritional interventions, as well as the long-term impact of early malnutrition during therapy. It is our goal that this information can be used for future studies and to help develop evidence-based guidelines to modify existing treatment plans.

REFERENCES

- 1. Becker P, Nieman Carney L, Corkins, MR et al. Consensus statement of the Academy of Nutrition and Dietetics/American Society for Parenteral and Enteral Nutrition: indicators recommended for the identification and documentation of pediatric malnutrition (undernutrition). J Acad Nutr Diet.
- 2. Brinksma A, Sanderman R, Roodbol, PF et al. Malnutrition is associated with worse health-related quality of life in children with cancer. Support Care Cancer. 2015;23(10):3043-3052.
- 3. Centers for Disease Control and Prevention. Z-score data files. CDC growth charts.
- www.cdc.gov/growthcharts/zscore.htm
- 4. Esbenshade AJ, Simmons JH, Koyama T, Koehler E, Whitlock JA, Friedman DL. Body mass index and blood pressure changes over the course of treatment of pediatric acute lymphoblastic leukemia. Pediatr Blood Cancer. 2011;56(3):372-378.
- 5. Fang Zhang F, Mae Rodday A, Kelly MJ, et al. Predictors of Being Overweight or Obese in Survivors of Pediatric Acute Lymphoblastic Leukemia (ALL).
- 6. Loeffen EAH, Brinksma A, Tissing WJE. Clinical implications of malnutrition in children with cancer.

 Support Care Cancer. 2015.

 Notice of Care and Partitute. Bigle Crowns for Childhead Acute Lymphablactic Levels and Childhead Acute.
- National Cancer Institute. Risk Groups for Childhood Acute Lymphoblastic Leukemia. Childhood Acute Lymphoblastic Leukemia Treatment.
- 8. Orgel E, Sposto R, Malvar, J et al. Impact on survival and toxicity by duration of weight extremes during treatment for pediatric acute lymphoblastic leukemia: a report from the Children's Oncology Group. J Clin Oncol. 2014;32(13):1331-1337.
- 9. Owens JL, Hanson SJ, McArthur JA, Mikhailov TA. The need for evidence based nutritional guidelines for pediatric acute lymphoblastic leukemia patients: Acute and long-term following treatment. Nutrients. 2013. 10. Winkler MR, Hockenberry MJ, McCarthy KS, Silva SG. Trajectories of Obesity and Overweight Rates
- Among Survivors of Childhood Acute Lymphoblastic Leukemia. Oncol Nurs Forum. 2015;42(4):336. 11.Withycombe J, Post-White J, Meza, JL et al. Weight patterns in children with higher risk ALL: A report from the Children's Oncology Group (COG) for CCG 1961. Pediatr Blood Cancer2. 2009;53(7):1249-
- 12.Zhang FF, Kelly MJ, Saltzman E, Must A, Roberts SB, Parsons SK. Obesity in Pediatric ALL Survivors: A Meta-Analysis. Pediatrics. 2014.
- 13.den Hoed MAH, Pluijm SMF, de Groot-Kruseman HA, et al. The negative impact of being underweight and weight loss on survival of children with acute lymphoblastic leukemia. Haematologica. 2015.

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